(19) World Intellectual Property Organization

International Bureau



A TABAN BUMAN KECININ BUMAN KANTAN BUMAN BANTAN BANTAN BANTAN BANTAN BANTAN BANTAN BANTAN BUMAN BANTAN BANTAN B

(43) International Publication Date 8 July 2004 (08.07.2004)

(10) International Publication Number WO 2004/056198 A1

(51) International Patent Classification7: A23L 1/0522, 1/0534

A23K 1/18,

(74) Agents: CORNISH, Kristina, Victoria, Joy et al.; Kil-

(21) International Application Number:

PCT/GB2002/005913

(22) International Filing Date:

23 December 2002 (23.12.2002)

(25) Filing Language:

English

(26) Publication Language:

English

- (71) Applicant (for all designated States except US): MARS UK LIMITED [GB/GB]; 3D Dundee Road, Slough, Berkshire SL1 4LG (GB).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): BUTTERWICK, Richard [GB/GB]; Waltham Centre for Pet Nutrition, Freeby Lane, Waltham-on-the-Wolds, Leicestershire LE14 4RT (GB). ROLFE, Vivien [GB/GB]; Waltham Centre for Pet Nutrition, Freeby Lane, Waltham-on-the-Wolds, Leicestershire LE14 4RT (GB).

- burn & Strode, 20 Red Lion Street, London WC1R 4PJ (GB).
- (81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
- (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

with international search report

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: FOOD FOR GASTROINTESTINAL HEALTH

(57) Abstract: The present invention relates to a foodstuff comprising a source of rice starch, a non-fermentable fibre and a bulk forming fermentable fibre and, in particular, its use in improving or maintaining the gastrointestinal health of a dog.

10

15

20

25

Food For Gastrointestinal Health

The present invention relates to a foodstuff comprising a source of rice starch, a non-fermentable fibre and a bulk forming fermentable fibre and its use in improving or maintaining the gastrointestinal health of a dog. The invention further relates to a method of improving the gastrointestinal health of a dog.

It has been observed that a proportion of the dog population exhibit non-specific dietary sensitivity on a range of foodstuffs. This dietary sensitivity can manifest as a variety of clinical symptoms such as vomiting, diarrhoea, skin disease, respiratory disorders and disorders of the central nervous system. The causes or dietary drivers of this dietary sensitivity remain elusive. However, common reported allergens in dogs include cow's milk, beef, mutton, pork, chicken, rabbit, horse, some fish, eggs, oatmeal, corn, wheat, soy, rice, potatoes and kidney beans.

Non-specific diet sensitivity is observed with all diets, however it is particularly associated with dogs fed on wet (moisture of 70 to 90%) or semi-wet (moisture of 15 to 70%) foodstuffs. The conventional solution to such non-specific diet sensitivity is to provide the dog with a "sensitive" product usually in the form of a dry diet. However, such a solution is not appropriate or desirable for all dogs. In addition, while a wet diet may exacerbate non-specific dietary sensitivity, sensitivity can also be associated with dry diets. Non-specific dietary sensitivity involves factors such as stress, activity levels and dietary components.

10

15

20

25

The present invention provides a foodstuff, in particular a wet or semi-wet foodstuff, which can be used to improve and/or treat the symptoms of canine non-specific dietary sensitivity. This foodstuff will allow dogs suffering from non-specific dietary sensitivity to be fed on a wet or semi-wet diet. This will avoid the problems associated with changing a dogs diet from a wet to a dry diet. In addition, the provision of a wet or semi-wet foodstuff provides more choice and flexibility to the owner.

A first aspect of the present invention provides a foodstuff comprising a source of rice starch, a source of non-fermentable fibre and a source of bulk forming fermentable fibre. For the purposes of this invention, the foodstuff may have a moisture content of from 15 to 90% and is preferably wet (moisture content of 70 to 90%) or semi wet (moisture content of from 15 to 70%).

The foodstuff of the first aspect contains a source of rice starch. The source of the rice starch is not limiting. It can be provided, for example, as rice (either whole or broken grains), ground rice or rice flour. The foodstuff further provides a source of non-fermentable fibre. The source of non-fermentable fibre is not limiting. It may be one or more of cellulose, wheat bran, oat bran or barley bran. The foodstuff further contains a source of a bulk forming fermentable fibre. For the purpose of this invention, bulk forming fibres improve faecal bulk thereby improving transit and laxation. The source of the bulk forming fibre is not limiting. Preferably, the bulk forming fermentable fibre is one or more of sugar beet pulp, coconut endosperm fibre, chicory pulp, citrus pulp, carob bean or gum talha.

10

15

20

25

In a preferred feature of the invention, a foodstuff is provided comprising rice starch, sugar beet pulp (as a source of bulk forming fermentable fibre) and cellulose (as a source of non-fermentable fibre). The sugar beet pulp is provided at a level of approximately 5% to approximately 0.1% weight/dry weight, preferably, approximately 3% to approximately 0.5%, more preferably at a level of approximately 1.6% or above. Cellulose is provided at a level of 5% to 0.1% weight/dry weight, preferably, approximately 2% to approximately 0.5%, more preferably at a level of approximately 0.8% or above. Rice starch is provided at a level of approximately 5% to approximately 0.1% weight/dry weight, preferably approximately 5% to approximately 0.1% weight/dry weight, preferably approximately 3% to approximately 0.5%, more preferably at a level of approximately 1.6% or above.

The levels of fibre in a foodstuff can be analysed using the Englyst method (as defined in Englyst H.N., and Cumming J.H. (1984), Simplified method for the measurement of total non-starch polysaccharides by gas-liquid chromatography of constituent sugars as alditol acetates. Analyst. 109, 937-942, and incorporated herein by reference). A description of the Englyst method is described in Appendix 1. In principle, starch is removed enzymatically after solubilisation and NSP is measured as the sum of the constituent sugars released by acid hydrolysis. The starch component of the fibre source is gelatinised by boiling in hot water and is then removed with alph-amylase and pullulanase. Starch and modified, or resistant starch are dispersed in DMSO. Three samples are then subjected to complementary procedures measuring (I) total NSP (ii) watersoluble NSP and (iii) cellulose. Components are hydrolysed in each case with The constituent sugars are converted to alditols and are sulphuric acid. measured as their alditol acetates using gas-liquid chromatography (GLC). Values for total dietary fibre as well as insoluble and soluble fractions can be

obtained. Cellulose can be measured separately and the non-cellulose polysaccharides are characterised by measurement of the individual monosaccharides.

The level of fibre in any particular fibre source can be determined by identifying the amount of fibre therein and comparing the level of fibre provided with that provided by the cellulose or sugar beet pulp as discussed above. The amount of a particular fibre source to add to a foodstuff can then be determined.

10

15

5

When the foodstuff of the first aspect of the invention is provided as a snack or treat, the levels of bulk forming fermentable fibre, non-fermentable fibre and rice starch can be decreased. For example, a snack food may be provided with approximately 0.8% rice starch or above, approximately 0.4% cellulose or above and approximately 0.8% sugar beet pulp or above.

In a preferred feature of the first aspect, the combined levels of non-fermentable fibre and bulk forming fermentable fibre does not exceed approximately 8% w/w, preferably the combined level does not exceed approximately 5% w/w.

20

25

The foodstuff according to the present invention encompasses any product that a pet consumes in its diet. Thus, the invention covers standard food products as well as pet food snacks (for example, snack bars, biscuits and sweet products – Preferably, these snackfoods are wet or semi-wet products such as co-extruded pet treats described in EP0647410 or WO99/47000. The foodstuff is preferably a cooked product. It may incorporate meat or animal derived material (such as beef, chicken, turkey, lamb, fish, blood plasma, marrow bone etc or one or more

thereof). The product alternatively may be meat free (preferably including a meat substitute such as soya, maize gluten or a soya product) in order to provide a protein source. The product may contain additional protein sources such as soya protein concentrate, milk proteins, gluten etc. The product may also contain an additional starch source (in addition to the source of rice starch) such as one or more grains (e.g. corn, rice, oats, barley etc).

The foodstuff of the present invention may preferable be provided as a liquid supplement. The supplement may be provided as an accompaniment with food or may be added to a conventional foodstuff. Alternatively, the supplement may be provided before or after the conventional foodstuff. The supplement may further be added to a drink such as milk or water.

The foodstuff is preferably packaged. In this way, the consumer is able to identify, from the packaging, the ingredients in the foodstuff or food supplement and confirm that it is suitable for the particular pet in question. The packaging may be metal (usually in the form of a tin or flexifoil), plastic, paper or card. The amount of moisture in any product may influence the type of packaging, which can be used or is required.

20

25

5

10

15

According to the present invention, dogs are any canine animal, in particular the domestic or pet dog, *Canis domesticus*.

The second aspect of the invention relates to a process for the manufacture of a foodstuff of the first aspect of the invention. The process comprises admixing a source of rice starch, a non-fermentable fibre and a bulk forming fermentable fibre. The foodstuff can be made according to any method known the art such

20

25

as in Waltham Book of Dog and Cat Nutrition, Ed. ATB Edney, Chapter by A. Rainbird, entitled "A Balanced Diet" in pages 57 to 74 Pergamon Press Oxford. All preferred features of the first aspect also apply to the second.

The third aspect of the invention relates to a foodstuff of the first aspect for use in improving and/or maintaining the gastrointestinal health of a dog. Improving and/or maintaining the gastrointestinal health of an animal is a long held aim in the art, particularly in dogs suffering from non-specific dietary sensitivity. The ability to maintain and/or improve gastrointestinal tract health can be beneficial to pet owners because it has an impact on their pet's overall health.

A dog with non-specific dietary sensitivity has sub-optimal intestinal health. This increases the risk of the dog developing viral or bacterial infections and compromises its long-term health. The foodstuff of the invention is preferably provided for improving and/or maintaining the gastrointestinal health of a dog with a canine non-specific dietary sensitivity.

The inventors have previously showed that dogs with a non-specific dietary sensitivity exhibit impaired water and electrolyte absorption. Furthermore, a dog with non-specific dietary sensitivity also exhibits a rapid whole gut transit time. These colonic abnormalities result in poor faeces. In addition, dogs with non-specific dietary sensitivity have diarrhoea and sub-optimal intestinal health. Without being bound by scientific theory, the foodstuff of the first aspect is believed to drive absorption and regulate whole gut transit time in dogs with non-specific dietary sensitivity. This leads to an improvement in the gastrointestinal health of these dogs.

By improving the gastrointestinal health of the animal, the invention seeks to promote and maintain good quality faeces in pet animals. Good faeces quality is of two-fold importance. Firstly, it is a good indicator of a healthy pet. It is known that good faeces quality usually reflects healthy colonic structure and function. Secondly, it is a much-favoured practicality for pet-owners. The invention therefore provides a foodstuff of the first aspect for improving and/or maintaining faeces quality in a dog.

Improving and/or maintaining gut health includes: improving and/or maintaining the gut motility of a dog. The foodstuff of the first aspect improves whole gut transit time in a dog with non-specific dietary sensitivity; improving and/or maintaining the absorption of electrolytes and colonic water in the gastrointestinal tract of a dog. This improves faeces quality and prevents and/or reduces diarrhoea in a dog with non-specific dietary sensitivity.

15

10

5

By improving the gastrointestinal health of a dog, the gastrointestinal tract is able to operate more efficiently, leading to further improvements in the overall health of the dog.

It has been found that the foodstuff of the first aspect of the invention comprising a source of rice starch, a source of non-fermentable fibre and a source of bulk forming fermentable fibre is more efficient and effective than a foodstuff containing one or a combination of two of the components. It is therefore submitted that the components, a source of rice starch, a source of non-fermentable fibre and a source of bulk forming fermentable fibre interact to provide a synergistic result. The foodstuff therefore provides improved benefits to a dog with non-specific dietary sensitivity.

10

15

20

The foodstuff of the third aspect can be administered to a dog in place of its conventional food. The foodstuff can be administered alone or in combination with a dry food or snack. Preferably, the foodstuff of the invention is administered to the dog daily, more preferably twice daily. Where the foodstuff is administered as a snack or treat, the foodstuff is administered to the dog one or more times a day, for example up to five times daily.

The fourth aspect of the invention relates to the use of a foodstuff of the first aspect to improve and/or maintain the gastrointestinal health of a dog. The dog is preferably in need of improvement in gastrointestinal health and may suffer from non-specific dietary sensitivity.

All preferred features of the first, second and third aspects of the invention also relate to the fourth aspect.

The fifth aspect of the invention relates to a method of improving the gastrointestinal health of a dog. The method comprises administering to the dog, the foodstuff of the first aspect. The dog may be suffering from non-specific dietary sensitivity. Administration is preferably by feeding.

All preferred features of the first, second, third and fourth aspects also apply to the fifth aspect.

The sixth aspect of the invention relates to the use of a source of rice starch, a non-fermentable fibre, and a bulk forming fibre in the manufacture of a

foodstuff wherein the foodstuff is for improving and/or maintaining gastrointestinal health in a dog.

All preferred features of the first, second, third, fourth and fifth aspects of the invention also apply to the sixth aspect.

The invention will now be illustrated with reference to the following nonlimiting examples.

10 **EXAMPLES**

Methods

15

20

25

A panel of ten control and ten sensitive dogs were fed either a standard foodstuff or a foodstuff comprising rice starch, cellulose and sugar beet pulp (supplemented food). The dogs were fed in accordance with individual energy requirements.

Both foodstuffs were wet foods with moisture levels of 78-79%. The major protein sources for each diet were poultry, beef, wheat and maize. The supplemented food was supplemented at the levels set out below (percentages relate to weight per dry weight):

Rice starch 1.6%

Cellulose 0.8%

Sugar beet pulp 1.6%

PCT/GB2002/005913

The diets were fed in a cross-over design that included a washout phase.

Experimental Design Summary				
Timings	Control (n=5)	Control (n=5)	Sensitive (n=5)	Sensitive (n=5)
Phase 1 3 weeks	Std	Std	Std	Std
1 week	Std	Supplemented food	Std	Supplemented food
Phase 2	Std Washout		Std Washout	
Phase 3 3 weeks	Std	Std	Std	Std
1 week	Supplemented food	Std	Supplemented food	Std

Measurements

5

10

Faeces quality was assessed daily using the WCPN 17-point linear scale. All defecations were scored with a score of grade 1 representing dry crumbly faeces and grade 5 diarrhoea. Major intermediate points are at grade 2 - ideal, well formed, does not leave a mark, easy to pick up; grade 3 - good quality slightly moist, less well formed, leaves a marked when removed from a dry surface, tacky to the touch, soft centred; and grade 4 – poor quality, moist, badly formed faeces with consistency of putty or porridge. Statistical comparisons were made on the 7-day period during which supplemented food was fed.

At the end of phase 1 and 3 standard and functional measurements were made: 15

Colonic transport function was measured using dialysis bags.

Whole gut transit time was measured by calculating the mean rate of transit of

barium impregnated polyethylene spheres (BIPS) through the intestinal tract. Briefly, 20 pellets were administered with the early morning feed and all faeces voided collected for 4 days. Faeces were x-rayed and the numbers of BIPS evacuated counted.

5

The mean whole gut transit time (WGTT) was calculated using the following equation:

WGTT (hours)
$$\sum$$
 no. pellets (I) x time interval (i) $_{(i=1-n)}$ \sum no. pellets (I) $_{(i=1-n)}$

10 Statistics

All data is expressed as the mean \pm the standard error of the mean. Statistical significance of all parameters was measured using Multifactoral Anova with significance assumed at p<0.05.



RESULTS

	Faeces Qua	ality Results	
	Mean Score (SEM)	% Unacceptable	Faecal output (per dog per day)
Control			
Std	2.3 (0.05) a	2.4	1.7 a
Supplemented Food	2.1 (0.03) a	0.5	1.9 a
Sensitive			
Std	2.7 (0.05) c	11.1	2.5 a
Supplemented Food	2.4 (0.04) b	3.9	2.3 a

The feeding of diet supplemented with rice starch, cellulose and sugar beet dramatically improved the faeces quality of the sensitive dogs.

Colonic Electrolyte Transport

Electrolyte transport (mM.hour) (-ve value = absorption)		
	Na+	
Control		
Std	-53.3 (10) b	
Supplemented food	-65.8 (8.8) b	
Sensitive		
Std	-64.1 (9.4) b	
Supplemented food	-94.9 (9.3) a	

Sensitive dogs showed significantly improved sodium absorption (indicating by a more negative value) from the colon when fed the supplemented foodstuff.

Whole Gut Transit Time

Administration of the supplemented food to sensitive dogs significantly improved the whole gut transit time, as indicated by an increase in the time of transit, improving times present to those observed in controlled dogs.

Mean Whole Gut Transit Time (Hours) [Standard Deviation]		
Control dogs		
Standard food	31.1 [10.6]b	
Supplemented food	26.0 [5.2]ab	
Sensitive dogs		
Standard food	21.6 [5.2]a	
Supplemented food	28.4 [8.5]b	

Effectiveness of foods comprising rice starch, cellulose or sugar beet pulp on faeces quality.

	Standard food	0.8% Cellulose	1.6% Sugar beet pulp	1.6% Rice starch
Control dogs	2.28 [†] 0.4 b	2.13 [†] 0.23 a	2.23 ⁺ 0.37 b	2.23 ⁺ 0.40 b
Sensitive dogs	2.28 ⁺ 0.39 b	2.15 ⁺ 0.24 a	2.21 ⁺ 0.28 ab	2.23 ⁺ 0.34 ab

Thus, the effects of feeding each of the components separately provides less benefit than providing a combination of the components.

10



Appendix 1

The Englyst method, from Englyst and Cummings (Supra).

5 Experimental

Apparatus

The fractionation procedure was carried out in 50-60ml screw-topped glass centrifuge tubes as previously described. Gas-liquid chromatography was performed with a Pye Unicam Series 204 chromatograph, fitted with a flame-ionisation detector. A 2.1m x 2mm i.d. glass column packed with Supelcoport (100-200 mesh) coated with 3% SP 2330 was used. The column temperature was 215°C (isothermal) and the injector and detector temperatures were 250°C. The carrier gas (nitrogen) flow-rate was 20ml min-1.

15

10

Reagents

High purity certified reagents were used for all analyses. Enzyme preparations were as follows: hog pancreatic α-amylase, E.C.3.2.1.1. (Sigma, Cat. No. A4268); pullulanase, E.C.3.2.1.41. (Boehringer, Cat. No. 108944).

20

Method

The sequence of steps in the procedure is summarised below.

Pre-treatment of sample

As far as possible, foods should be analysed without any pre-treatment. If there are problems in taking a representative sample, foods with a low water content

can be ball milled for 2-3 minutes, and those with a higher water content homogenised, or freeze-dried and ball milled.

Sample Mass

Accurately weigh between 50 and 1,000mg of sample, containing not more than 150mg of starch and 50mg of NSP, into a 50-60ml screw-top centrifuge tube and add a stirrer.

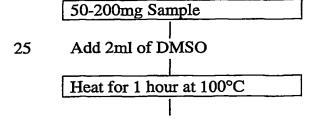
Fat Extraction and Drying

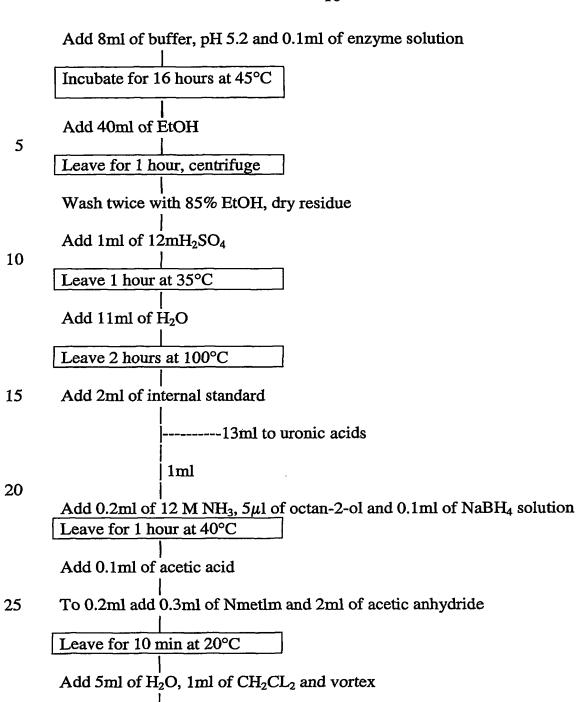
Samples with dry matter between 90 and 100% and with less than 203% of fat can be analysed directly. Otherwise, add 40ml of acetone, mix for 30 minutes by using a magnetic stirrer, centrifuge and remove by aspiration as much of the supernatant as possible without disturbing the residue. Place the tubes in a water bath at 65°C on a magnetic stirrer hot plate and mix the residue for a few minutes until it appears to be dry. The beaker can be covered and the acetone vapour removed by water pump.

Dispersion of the Starch

20

Add 2ml of DMSO, cap the tube and heat it in a boiling water bath for 1 hour, timed from when re-boiling commences, stirring continuously. Then, without cooling, add 8ml of 0.1M sodium acetate buffer pH5.2, at 50°C and vortex mix immediately.





Procedure for the analysis of non-starch polysaccharides (NSP)

Use 1-2 μ l of the lower phase for GLC

30

10

15

20

25

Enzyme Hydrolysis of the Starch

Cool the tube to 45°C and immediately add 0.1ml of an enzyme solution containing 5,000 units of α-amylase and 5 units of pullulanase per ml of acetate buffer at pH 5.2. Incubate the samples at 45°C for 16-18 hours, preferably mixing continuously as described previously.

Following the enzyme treatment, add 40ml of absolute ethanol, mix well and leave to stand for 1 hour at room temperature. Centrifuge for 10 minutes or until a clear supernatant liquid is obtained. Removed by aspiration as much of the supernatant liquid as possible, without disturbing the residue, and discard it. Wash the residue twice with 50ml of 85% ethanol by mixing to form a suspension, centrifuging until clear and removing the supernatant liquid as before. Add 40ml of acetone to the washed residue, stir for 5 minutes and then centrifuge. Remove the supernatant liquid by aspiration and dry the residue as described under *Fat extraction and drying*.

Acid hydrolysis of the residue from enzymic digestion

Disperse the dried residue in 1ml of 12M sulphuric acid, using a vortex mixer. Leave at 35°C for 1 hour to solubilise the cellulose, then rapidly add 11ml of water and mix.

Heat the solution in a boiling water bath for 2 hours from re-boiling, stirring continuously. Cool it to room temperature by placing the tube in water, add 2ml of internal standard (2 mg of allose per ml of saturated benzoic acid solution) and mix the contents of the tube. Use 1ml of the hydrolysate for the

preparation of alditol acetates and keep the remainder for the determination of uronic acids.

Uronic acids

5

10

15

The method used is a modification of the method of Scott. Mix 0.3ml of hydrolysate (diluted, if necessary, so that it contains between 25 and $100\mu\text{g}$ of uronic acids per ml) with 0.3ml of a mixtures of sodium chloride-boric acid solution (prepared by adding 2g of sodium chloride and 3g of boric acid to 100ml of water) Add 5ml of concentrated sulphuric acid and vortex mix, then place the tube in a heating block at 70°C . Leave the tube and contents for 40 minutes and then cool them to room temperature by placing in water. When cool, add 0.2ml of 3.5-dimethylphenol solution (0.1g of $(\text{CH}_3)_2\text{-C}_6\text{H}_3\text{OH}$ in 100ml of glacial acetic acid) and mix immediately. Between 10 and 15 minutes later read the absorbance at 400 and 450nm in a spectrophotometer against a water reference. Subtract the reading at 400nm from that at 450nm for each sample and plot the difference obtained for glucuronic acid standards (over the range $25\text{-}125\mu\text{f} \text{ ml}^{-1}$). Read the sample concentrations from the graph.

Preparation of alditol acetates

To 1ml of hydrolysate add 0.2ml of 12M ammonia solution and 5μl of octan-2ol. Test that the solution is alkaline, and then add 0.1ml of a freshly prepared solution of 100mg of sodium tetrahydroborate (III) (sodium borohydride) per ml of 3M ammonia solution. Mix, leave the mixture for 1 hour at 40°C and add 0.1ml of glacial acetic acid. Next, to 0.2ml of the acidified solution add 0.3ml of N-methylimidazole and 2ml of acetic anhydride, and mix. Leave it for 10 minutes at 20°C (room temperature), add 5ml of water, mix, and when cooled add 1ml of dichloromethane, agitate the contents vigorously on a vortex mixer and centrifuge for a few minutes to separate the mixture into two phases. Remove the bulk of the upper phase by aspiration and discard it, then transfer the lower phase to a small vial, seal and store it at -20° C. Use $1-2\mu$ l for injection on to the chromatograph.

5

10

15

20

Alternative preparative of alditol acetates

When dichloromethane is used as a solvent for the alditol acetates it has been observed in a number of laboratories without automatic GLC injection facilities that the injection technique is critical to the obtaining of reproducible results. A more robust method can be obtained if dichloromethane is replaced with ethyl acetate as a solvent for alditol acetates. The procedure is as follows:

To 1ml of hydrolysate add 0.2ml of 12M ammonia solution and 5μ l of octan-2-ol. Test that the solution is alkaline, then add 0.1ml of a freshly prepared solution of 100mg of sodium tetrahydroborate (III) per ml of 3M ammonia solution. Mix, leave the mixture for 1 hour at 40°C and add 0.1ml of glacial acetic acid. To 0.5ml of the acidified solution add 0.5ml of N-methylimidazole, 5ml of acetic anhydride and mix. Leave for 10 minutes at 20°C (room temperature), then add 0.6ml of ethanol and mix. After 5 minutes add 5ml of water, place in a water bath at room temperature, add 5ml of 7.5M KOH and a few minutes later a further 5ml of 7.5M KOH. Mix by inverting and leave to separate into two phases. Transfer the top phase to a small vial and store at +5°C. Use $1-2\mu$ l for injection on the chromatograph.

CLAIMS

5

10

25

- 1. A foodstuff comprising a source of a rice starch, a non-fermentable fibre and a bulk forming fermentable fibre.
- 2. A foodstuff as claimed in claim 1 which is a wet or semi-wet foodstuff.

20

- 3. A foodstuff as claimed in claim 1 or claim 2 wherein the source of rice starch is rice, ground rice or rice flour.
- 4. A foodstuff as claimed in any one of claims 1 to 3 wherein the non-fermentable fibre is cellulose.
- 5. A foodstuff as claimed in any one of claims 1 to 4 wherein the bulk forming fermentable fibre is sugar beet pulp.
 - 6. A foodstuff as claimed in any one of claims 1 to 5 for use in improving the gastrointestinal health of a dog.
- 7. A foodstuff as claimed in claim 6 wherein the dog has non-specific dietary sensitivity.
 - 8. A foodstuff as claimed in claim 6 for use in improving intestinal function in a dog.
 - 9. A foodstuff as claimed in claim 6 for use in improving gut motility in a dog.

WO 2004/056198

10

- 10. A use of a foodstuff as claimed in any one of claims 1 to 5 to improve the gastrointestinal health of a dog.
- 5 11. A method of improving the gastrointestinal health of a dog comprising administering a foodstuff as claimed in claim 11.
 - 12. A method for preventing diarrhoea in a dog comprising administering a foodstuff as claimed in any one of claims 1 to 5.
 - 13. The use of rice starch, a non-fermentable fibre and a bulk forming fermentable fibre in the manufacture of a foodstuff for improving the gastrointestinal health of a dog.
- 14. A process for the manufacture of a foodstuff as claimed in any one of claims 1 to 9 comprising admixing a source of rice starch, a non-fermentable fibre and a bulk forming fermentable fibre.
- 15. A foodstuff as claimed in claims 1 to 9 comprising of rice starch at a level of approximately 5% to approximately 0.1% w/w.
 - 16. A foodstuff as claimed in claims 1 to 9 comprising sugar beet pulp at a level of approximately 5% to approximately 0.1% w/w.
- 25 17. A foodstuff as claimed in any one of claims 1 to 9 comprising cellulose at a level of approximately 5% to approximately 0.1% w/w.

A. CLASSIFICATION OF SUBJECT MATTER
IPC 7 A23K1/18 A23L1/0522 A23L1/0534

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, BIOSIS, MEDLINE, FSTA

	ENTS CONSIDERED TO BE RELEVANT	Delevent to eleim No.
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Χ	WO 98 44932 A (IAMS COMPANY)	1-6,8,
	15 October 1998 (1998-10-15)	10,11, 15-17
	page 2-3	
	tables 1,3	
	claims 1,7,9,15	,
_	DINAME COMMINE MICHAEL MICHAEL	1-12,
ı	DINAND E, CHANZY H, VIGNON MR:	15-17
	"Suspensions of cellulose microfibrils from sugar beet pulp"	15 17
	FOOD HYDROCOLLOIDS,	
	vol. 13, 1999, pages 275-283, XP002251873	
	page 275, column 1, paragraph 1	
	page 277, column 2, paragraph 3	
	_/	
ſ		
ı		

Further documents are listed in the continuation of box C.	χ Patent family members are listed in annex.
Special categories of cited documents: A' document defining the general state of the art which is not considered to be of particular relevance E' earlier document but published on or after the international filing date L' document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) O' document referring to an oral disclosure, use, exhibition or other means P' document published prior to the international filing date but later than the priority date claimed	 "T" later document published after the International filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. "&" document member of the same patent family
Date of the actual completion of the international search	Date of mailing of the international search report
27 August 2003	11/09/2003
Name and mailing address of the ISA	Authorized officer
European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31–70) 340–2040, Tx. 31 651 epo nl, Fax: (+31–70) 340–3016	Couzy, F

		P6 GB 02/05913
C.(Continua	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	WO 00 53030 A (BAILLON MARIE LOUISE AMANDA; MARS UK LTD (GB); FONE JANEL (GB); GI) 14 September 2000 (2000-09-14) page 3-7 page 9, line 22 page 10, line 11 page 11, line 23-25 page 13-15 examples 4,5,7 table 2 claims 1,2,12,13,17,20,21,24-28	1-17
A	ROLFE VE, ADAMS CA, BUTTERWICK RF, BATT RM: "Relationship between faecal character and intestinal transit time in normal dogs and diet-sensitive dogs" JOURNAL OF SMALL ANIMAL PRACTICE, vol. 43, 2002, pages 290-294, XP009015869 UK page 290, column 2, paragraphs 2,3 page 291, column 2, paragraph 3	1–17
	page 292, column 3 -page 293, column 1 page 294, column 1 figure 1; tables 1,2	1 17
A	WILLS J ET AL: "DIAGNOSIS AND MANAGEMENT OF FOOD ALLERGY AND INTOLERANCE IN DOGS AND CATS" AUSTRALIAN VETERINARY JOURNAL, AUSTRALIAN VETERINARY ASSOCIATION, BRUNSWICK, AU, vol. 71, no. 10, 1994, pages 322-326, XP008017281 ISSN: 0005-0423 page 325, column 1, paragraph 2	1-17
A	WG GUILFORD: "Nutritional management of gastrointestinal tract diseases of dogs and cats" JOURNAL OF NUTRITION, vol. 124, 1994, pages 2663S-2669S, XP009015863 USA page 2666S, column 1, paragraph 2 page 2666S, column 2, paragraph 3 -/	1-17

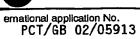


Int and Application No PCT GB 02/05913

	· · · · · · · · · · · · · · · · · · ·	•	05913
C.(Continua	Ition) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with Indication, where appropriate, of the relevant passages	R	elevant to claim No.
	SUNVOLD G D ET AL: "DIETARY FIBER FOR DOGS: IN VITRO FERMENTATION OF SELECTED FIBER SOURCES FOR DOG FECAL INOCULUM AND IN VIVO DIGESTION AND METABOLISM OF FIBER-SUPPLEMENT DIETS" JOURNAL OF ANIMAL SCIENCE, NEW YORK, NY, US, vol. 73, no. 4, 1995, pages 1099-1109, XP002072351 ISSN: 0021-8812 page 1101, column 2 -page 1102, column 1 page 1105, column 1-2 page 1106, column 1, paragraph 3 -column 2, paragraph 1 page 1108, column 2, paragraph 3		1-17



INTERNATIONAL SEARCH REPORT



Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This international Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely: see FURTHER INFORMATION sheet PCT/ISA/210
2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This international Searching Authority found multiple inventions in this international application, as follows:
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this international Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.



International Application No. PCT&B 02 \(05913 \)

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.1

Although claims 10-13 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.

Continuation of Box I.1

Rule 39.1(iv) PCT — Method for treatment of the human or animal body by therapy

INTERNATIONAL SEARCH REPORT

tion on patent family members

Int Phal Application No
PC 17 GB 02/05913

Patent document cited in search report	Publication date		Patent family member(s)	Publication date
WO 9844932 A	15-10-1998	AU AU	734098 B2 6796998 A	07-06-2001 30-10-1998
		BR	9807935 A	22-02-2000
		CN	1251995 T	03-05-2000
		EP	0967985 A1	05-01-2000
		JP	2000513583 T	17-10-2000
		NZ	337829 A	30-03-2001
		TR	9902485 T2	22-05-2000
		US	6180131 B1	30-01-2001
		WO	9844932 A1	15-10-1998
		US	2002197275 A1	26-12-2002
		US	6475512 B1	05-11-2002
		US	RE38112 E1	06-05-2003
WO 0053030 A	14-09-2000	AU	3175800 A	28-09-2000
WC CCCCCC	- · • · - · · ·	BR	0008879 A	22-01-2002
		CA	2366920 A1	14-09-2000
		CN	1345184 T	17-04-2002
		ΕP	1158867 A2	05-12-2001
		WO	0053030 A2	14-09-2000
		GB	2363696 A ,B	09-01-2002
		HU	0202350 A2	28-10-2002
		JP	2002537861 A	12-11-2002
		NZ	514054 A	28-09-2001
		PL	350537 A1	16-12-2002

This Page is Inserted by IFW Indexing and Scanning Operations and is not part of the Official Record

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images include but are not limited to the items checked:
☐ BLACK BORDERS
☐ IMAGE CUT OFF AT TOP, BOTTOM OR SIDES
FADED TEXT OR DRAWING
BLURRED OR ILLEGIBLE TEXT OR DRAWING
☐ SKEWED/SLANTED IMAGES
☐ COLOR OR BLACK AND WHITE PHOTOGRAPHS
☐ GRAY SCALE DOCUMENTS
LINES OR MARKS ON ORIGINAL DOCUMENT
☐ REFERENCE(S) OR EXHIBIT(S) SUBMITTED ARE POOR QUALITY
□ OTHER:

IMAGES ARE BEST AVAILABLE COPY.

As rescanning these documents will not correct the image problems checked, please do not report these problems to the IFW Image Problem Mailbox.